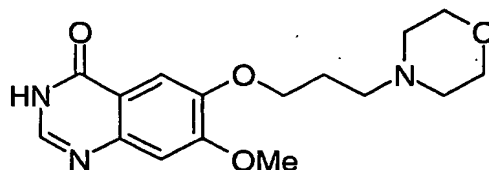


Claims

1. A process for the manufacture of 7-methoxy-6-(3-morpholinopropoxy)-3,4-dihydroquinazolin-4-one of Formula II

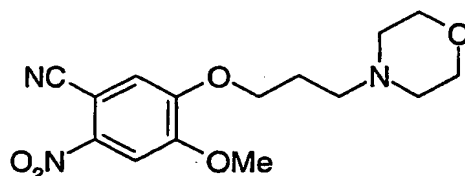


II

5

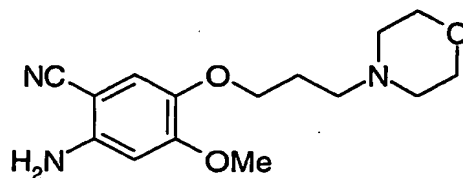
which comprises :-

- (a) the reduction of 4-methoxy-5-(3-morpholinopropoxy)-2-nitrobenzonitrile of Formula III



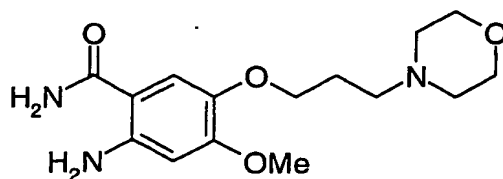
III

- 10 to give 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzonitrile of Formula IV



IV

- (b) the hydration of the compound of Formula IV to give 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzamide of Formula V

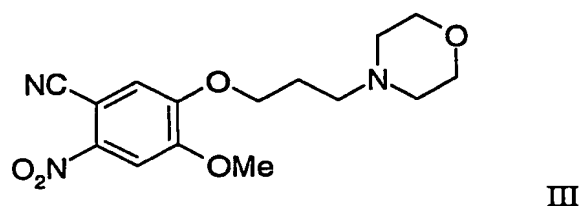


V

- 15 and (c) the cyclisation reaction of the compound of Formula V with formic acid, or a reactive derivative thereof, to give the compound of Formula II.

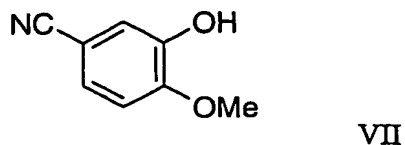
2. An intermediate of Formula III suitable for use in the process according to Claim 1.

3. An intermediate of Formula IV suitable for use in the process according to Claim 1.
4. A process according to Claim 1 wherein the intermediates of Formula IV and V are not
5 isolated as such but are each prepared and used as a solution in an organic solvent.
5. A process according to Claim 1 wherein step (a) is carried out in the presence of a
water-soluble inorganic reducing agent.
- 10 6. A process according to Claim 1 wherein step (a) is carried out by hydrogenation in the
presence of a suitable metal catalyst.
7. A process according to Claim 1 wherein step (b) is carried out in the presence of an
alkali metal base and in a polar protic solvent.
- 15 8. A process according to Claim 1 wherein step (c) is carried out in the presence of
formamide.
9. A process for the manufacture of 4-methoxy-5-(3-morpholinopropoxy)-
20 2-nitrobenzonitrile of Formula III

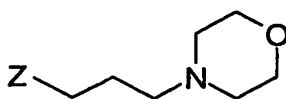


which comprises :-

- (a) the coupling of 3-hydroxy-4-methoxybenzonitrile of Formula VII

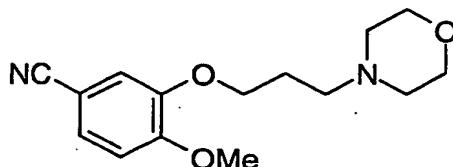


- 25 with a 3-morpholinopropane derivative of Formula VIII



VIII

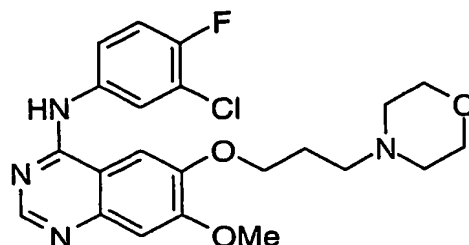
wherein Z is a displaceable group to give 4-methoxy-3-(3-morpholinopropoxy)benzonitrile of Formula IX



IX

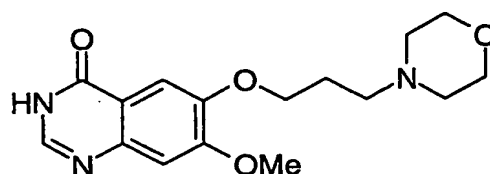
- 5 and (b) the nitration of the compound of Formula IX to give 4-methoxy-5-(3-morpholinopropoxy)-2-nitrobenzonitrile of Formula III.
10. An intermediate of Formula IX suitable for use in the process according to Claim 9.
- 10 11. A process according to Claim 9 wherein the intermediate of Formula IX is not isolated as such but is prepared and used as a solution in an organic solvent.
12. A process according to Claim 9 wherein the coupling reaction of step (a) is carried out as an alkylation reaction in the presence of an alkali metal carbonate.
- 15 13. A process according to Claim 9 wherein the coupling reaction of step (a) is carried out as a dehydration reaction in the presence of an azo compound and a phosphine.
14. A process according to Claim 9 wherein step (b) is carried out in the presence of nitric
- 20 acid.
15. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I

- 19 -



I

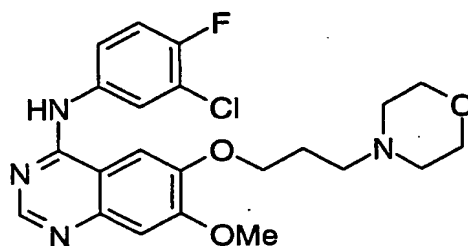
which comprises steps (a), (b) and (c) according to Claim 1 to manufacture 7-methoxy-6-(3-morpholinopropoxy)-3,4-dihydroquinazolin-4-one of Formula II



II

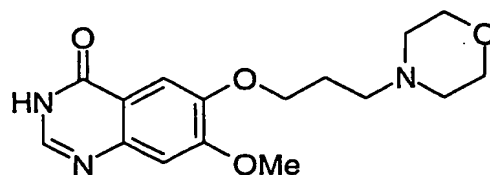
5 and the conversion of the compound of Formula II into the compound of Formula I.

16. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

10 which comprises steps (b) and (c) according to Claim 1 to manufacture 7-methoxy-6-(3-morpholinopropoxy)-3,4-dihydroquinazolin-4-one of Formula II

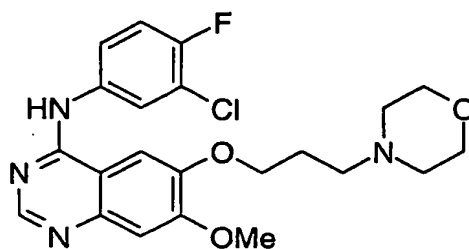


II

and the conversion of the compound of Formula II into the compound of Formula I.

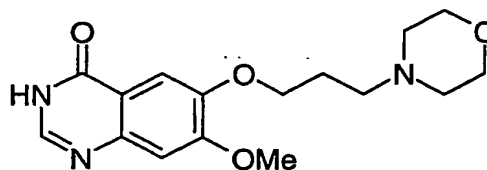
15 17. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-

6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

which comprises step (c) according to Claim 1 to manufacture 7-methoxy-6-(3-morpholinopropoxy)-3,4-dihydroquinazolin-4-one of Formula II

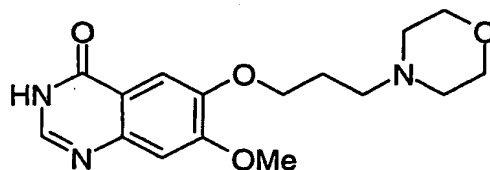


II

5

and the conversion of the compound of Formula II into the compound of Formula I.

18. The use of 7-methoxy-6-(3-morpholinopropoxy)-3,4-dihydroquinazolin-4-one of Formula II

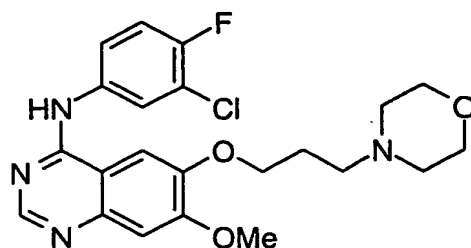


II

10

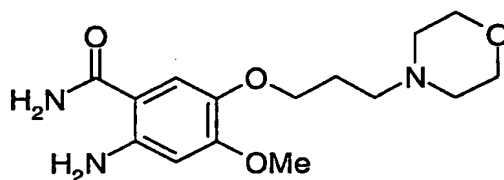
in the manufacture of a pharmacologically effective quinazoline derivative.

19. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

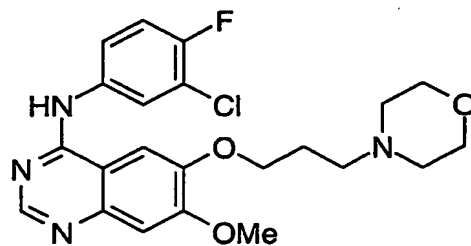
which comprises steps (a) and (b) according to Claim 1 to manufacture 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzamide of Formula V



V

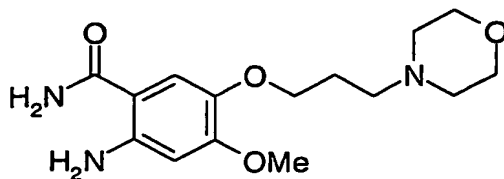
5 and the conversion of the compound of Formula V into the compound of Formula I.

20. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

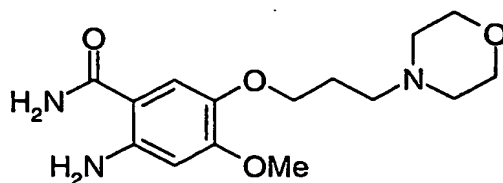
10 which comprises step (b) according to Claim 1 to manufacture 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzamide of Formula V



V

and the conversion of the compound of Formula V into the compound of Formula I.

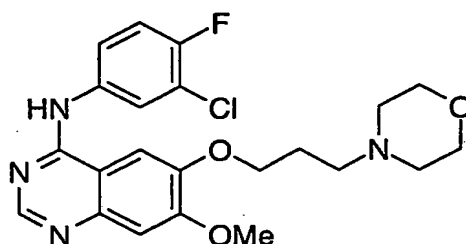
15 21. The use of 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzamide of Formula V



V

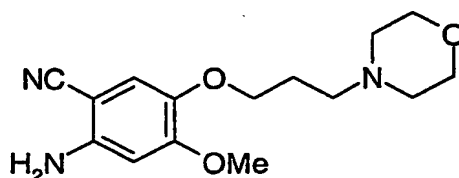
in the manufacture of a pharmacologically effective quinazoline derivative.

22. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-
5 6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

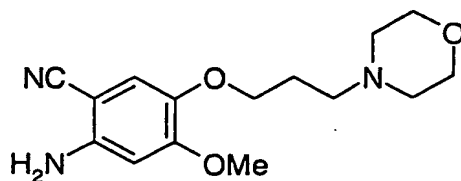
which comprises step (a) according to Claim 1 to manufacture 2-amino-4-methoxy-
5-(3-morpholinopropoxy)benzonitrile of Formula IV



IV

- 10 and the conversion of the compound of Formula IV into the compound of Formula I.

23. The use of 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzonitrile of Formula IV



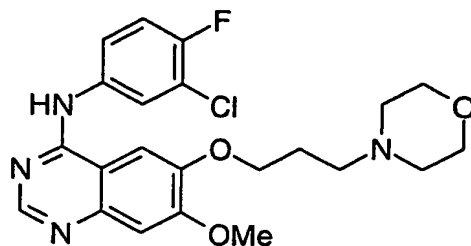
IV

in the manufacture of a pharmacologically effective quinazoline derivative.

15

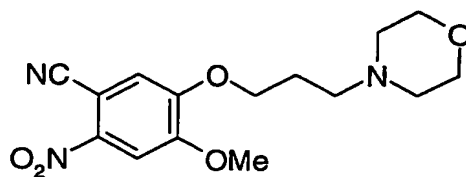
24. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-

6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

which comprises steps (a) and (b) according to Claim 9 to manufacture 4-methoxy-5-(3-morpholinopropoxy)-2-nitrobenzonitrile of Formula III

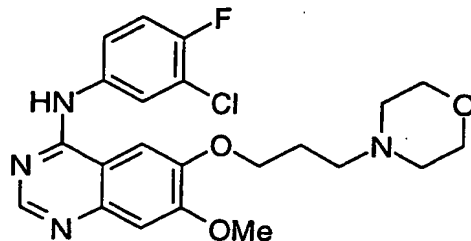


III

5

and the conversion of the compound of Formula III into the compound of Formula I.

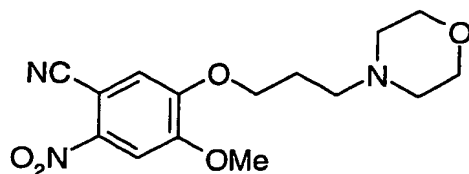
25. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

10

which comprises step (b) according to Claim 9 to manufacture 4-methoxy-5-(3-morpholinopropoxy)-2-nitrobenzonitrile of Formula III

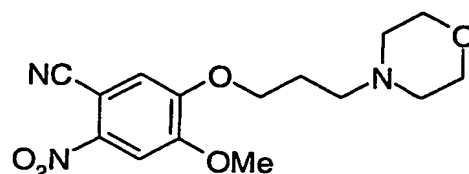


III

and the conversion of the compound of Formula III into the compound of Formula I.

15

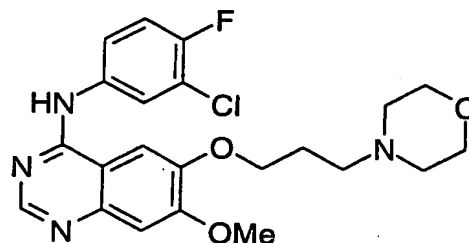
26. The use of 4-methoxy-5-(3-morpholinopropoxy)-2-nitrobenzonitrile of Formula III



III

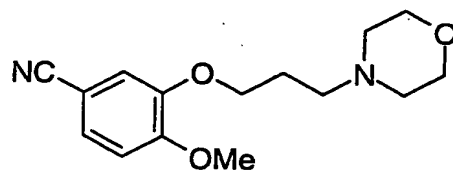
in the manufacture of a pharmacologically effective quinazoline derivative.

27. A process for the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I

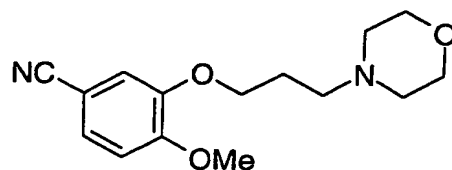
which comprises step (a) according to Claim 9 to manufacture 4-methoxy-3-(3-morpholinopropoxy)benzonitrile of Formula IX



IX

and the conversion of the compound of Formula IX into the compound of Formula I.

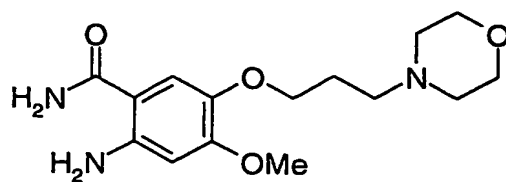
28. The use of 4-methoxy-3-(3-morpholinopropoxy)benzonitrile of Formula IX



IX

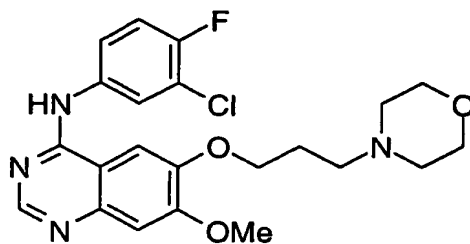
in the manufacture of a pharmacologically effective quinazoline derivative.

29. The use of 2-amino-4-methoxy-5-(3-morpholinopropoxy)benzamide of Formula V



V

in the manufacture of 4-(3'-chloro-4'-fluoroanilino)-7-methoxy-6-(3-morpholinopropoxy)quinazoline, the compound of Formula I



I